CLAIMS

A compound according to formula (I)

(I)

R1
$$C (CH2)n-A - (CH2)m-N (CH2)p O B$$

$$X$$

5 wherein:

is a phenyl ring, a C₄ to C₉ heteroaromatic compound containing one or more heteroatoms or a naphthalenyl, 5,6,7,8-tetrahydronaphthalenyl or biphenyl group;

 R^1 , R^2 and R^3 each independently represent a hydrogen atom or halogen atom, or a hydroxy group, or a phenyl, $-OR^4$, $-SR^4$, $-NR^4R^5$, $-NHCOR^4$, $-CONR^4R^5$, -CN, $-NO_2$, $-COOR^4$ or $-CF_3$ group, or a straight or branched lower alkyl group which may optionally be substituted, for example, with a hydroxy or alkoxy group, wherein R^4 and R^5 each independently represent a hydrogen atom,

15 straight or branched lower alkyl group or together form an alicyclic ring; or R^1 and R^2 together form an aromatic, alicyclic or heterocyclic ring,

n is an integer from 0 to 4;

A represents a $-CH_2-$, $-CH=CR^6-$, $-CR^6=CH-$, $-CR^6R^7-$, -CO-, -O-, -S-,

20 $-S(0)\text{-},\ SO_2$ or $-NR^6\text{-}$ group, wherein R^6 and R^7 each independently represent a hydrogen atom, straight or branched lower alkyl group or R^6 and R^7 together form an alicyclic ring;

m is an integer from 0 to 8; provided that when m = 0, A is not - CH_2 -;

p is an integer from 1 to 2 and the substitution in the azoniabicyclic ring may be in the 2, 3 or 4 position including

all possible configurations of the asymmetric carbons;
B represents a group of formula i) or ii):

(I)

wherein R^{10} represents a hydrogen atom, a hydroxy or methyl group; and R^{8} and R^{9} each independently represent

wherein R^{11} represents a hydrogen or halogen atom or a straight or branched lower alkyl group and Q represents a single bond, - CH_2 -, - CH_2 - CH_2 -, -O-, -O- CH_2 -, -S-, -S- CH_2 - or -CH=CH-; and X represents a pharmaceutically acceptable anion of a mono or polyvalent acid.

- 2. A compound according to claim 1, wherein any alkyl group present as R^1 to R^7 or R^{11} contains from 1 to 4 carbon atoms.
 - 3. A compound according to claim 1 or 2 wherein p=2.
- 4. A compound according to any one of the preceding claims wherein represents a phenyl, pyrrolyl, thienyl, furyl, biphenyl, naphthalenyl, 5,6,7,8-tetrahydronaphthalenyl, benzo[1,3]dioxolyl, imidazolyl or benzothiazolyl group.
- 5. A compound according to claim 4, wherein © represents a phenyl, pyrrolyl or thienyl group.

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6. A compound according to any one of the preceding

- claims wherein R^1 , R^2 and R^3 each independently represent a hydrogen or halogen atom or a hydroxy, methyl, tert-butyl, CH_2OH , 3-hydroxypropyl, -OMe, -NMe₂, -NHCOMe, -CONH₂, -CN, -NO₂, COOMe or -CF₃ group.
- 7. A compound according to claim 6 wherein R^1 , R^2 and R^3 each independently represent a hydrogen or halogen atom or a hydroxy group.
 - 8. A compound according to claim 7, wherein the halogen atom is fluorine.
- 9. A compound according to any one of the preceding claims wherein A represents a -CH₂-, -CH=CH-, -CO-, -NH-, -NMe-, -O- or -S- group; n is 0 or 1; and m is an integer from 1 to 6.
 - 10. A compound according to claim 9, wherein A represents a -CH2-, -CH=CH- or -O- group and m is 1, 2 or 3.
- 11. A compound according to any one of the preceding claims wherein the azoniabicyclo group is substituted on the nitrogen atom with a 3-phenoxypropyl, 2-phenoxyethyl, 3-phenylallyl, phenethyl, 3-phenylpropyl, 4-phenylbutyl, 3-(2-hydroxyphenoxy)propyl, 3-(4-fluorophenoxy)propyl, 2-
- 20 benzyloxyethyl, 3-pyrrol-1-ylpropyl, 2-thien-2-ylethyl or 3-thien-2-ylpropyl group.
 - 12. A compound according to any one of the preceding claims wherein B represents a group of formula (i) and R^8 and R^9 each independently represent a phenyl, 2-thienyl, 3-thienyl, 2-
- 25 furyl, or 3-furyl group and R¹¹ represents a hydrogen atom.
 - 13. A compound according to any one of claims 1 to 11, wherein B represents a group of formula (ii) and Q represents a single bond, a $-CH_2-$, $-CH_2-CH_2-$ group or an oxygen atom.
- 14. A compound according to any one of the preceding 30 claims wherein X represents a bromide, chloride or trifluoroacetate anion.
 - 15. A compound according to any one of the preceding claims wherein the azoniabicyclo group is substituted in the 3-

position.

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- 16. A compound according to claim 15, wherein the substituent in the 3 position has (R) configuration.
- 17. A compound according to claim 16, wherein R^8 is different from R^9 in group i), and the asymmetric carbon to which
- 18. A compound according to claim 16, wherein R^8 is different from R^9 in group i), and the asymmetric carbon to which R^8 and R^9 are bonded has the (S) configuration.
- 10 19. A compound according to any one of the preceding claims which is a single isomer.
 - 20. A compound according to claim 1 which is 3(R)-Diphenylacetoxy-1-(3-phenoxy-propyl)-1-azoniabicyclo[2.2.2]octane; bromide

 R^8 and R^9 are bonded has the (R) configuration.

- 3 (R) (2-Hydroxy-2,2-diphenyl-acetoxy) -1- (3-phenoxypropyl) -1azoniabicyclo[2.2.2]octane; bromide
 3 (R) (2,2-Diphenylpropionyloxy) -1- (3-phenoxypropyl) -1azoniabicyclo[2.2.2]octane; bromide
 - 3(R)-(2-Hydroxy-2-phenyl-2-thien-2-yl-acetoxy)-1-(3-
- phenoxypropyl)-1-azonia-bicyclo[2.2.2]octane; bromide
 3(R)-(2-Furan-2-yl-2-hydroxy-2-phenylacetoxy)-1-(3-phenylallyl)1-azo niabicyclo[2.2.2]octane; bromide
 - 3(R)-(2-Furan-2-yl-2-hydroxy-2-phenylacetoxy)-1-(2-phenoxyethyl)-1-azoniabicyclo[2.2.2]octane; bromide
- 25 3(R)-(2-Furan-2-yl-2-hydroxy-2-phenylacetoxy)-1-(3-phenoxypropyl)-1-azoniabicyclo[2.2.2]octane; bromide
 3(R)-(2,2-Dithien-2-ylacetoxy)-1-(3-phenoxypropyl)-1azoniabicyclo[2.2.2]octane; bromide
 3(R)-(2-Hydroxy-2,2-di-thien-2-ylacetoxy)-1-phenethyl-1-
- 30 azoniabicyclo[2.2.2]octane; bromide
 3(R)-(2-Hydroxy-2,2-di-thien-2-ylacetoxy)-1-(4-phenylbutyl)-1azoniabicyclo[2.2.2]octane; bromide
 3(R)-(2-Hydroxy-2,2-dithien-2-ylacetoxy)-1-(3-phenoxypropyl)-1-

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azonia-bicyclo[2.2.2]octane; bromide
    1-[3-(4-Fluorophenoxy)propyl]-3(R)-(2-hydroxy-2,2-dithien-2-ylace
    to xy)-1-azoniabicyclo[2.2.2]octane; chloride
    3(R)-(2-Hydroxy-2,2-dithien-2-ylacetoxy)-1-[3-(2-hydroxyphenoxy)p
5
   ro pyl]-1-azoniabicyclo[2.2.2]octane; trifluoroacetate
    3(R)-(2-Hydroxy-2,2-dithien-2-ylacetoxy)-1-(3-pyrrol-1-ylpropyl)-
    1-azonia-bicyclo[2.2.2]octane; trifluoroacetate
    3(R)-(2-Hydroxy-2,2-dithien-2-ylacetoxy)-1-(2-thien-2-ylethyl)-1-
    azo niabicyclo[2.2.2]octane; bromide
10
    3(R) - (2-Hydroxy-2, 2-dithien-2-ylacetoxy) - 1 - (3-thien-2-ylpropyl) - 1
    -a zoniabicyclo[2.2.2]octane; bromide
    1-(2-Benzyloxyethyl)-3(R)-(2-hydroxy-2,2-dithien-2-ylacetoxy)-1-a
    zoniabicyclo[2.2.2]octane; trifluoroacetate
    3(R)-(2-Hydroxy-2,2-dithien-3-ylacetoxy)-1-(3-phenoxypropyl)-1-az
15
    oniabicyclo[2.2.2]octane; bromide
    1-(3-phenylally1)-3(R)-(9-Hydroxy-9[H]-fluorene-9-carbonyloxy)-1-
    azoniabicyclo[2.2.2]octane; bromide
    3(R)-(9-Hydroxy-9[H]-fluorene-9-carbonyloxy)-1-(3-phenoxypropyl)-
    1-azoniabicyclo[2.2.2]octane; bromide
20
    3(R)-(9-Hydroxy-9[H]-fluorene-9-carbonyloxy)-1-phenethyl-1-azonia
    bicyclo[2.2.2]octane; bromide
    3(R)-(9-Hydroxy-9H-fluorene-9-carbonyloxy)-1-(3-thien-2-ylpropyl)
    -1-azoniabicyclo[2.2.2]octane; bromide
    3(R)-(9-Methyl-9[H]-fluorene-9-carbonyloxy)-1-(3-phenylallyl)-1-a
25
    zonia bicyclo[2.2.2]octane; bromide
    3(R)-(9-Methyl-9[H]-fluorene-9-carbonyloxy)-1-(3-phenoxypropyl)-1
    -azo niabicyclo[2.2.2]octane; bromide
    1-(4-Phenylbutyl)-3(R)-(9[H]-xanthene-9-carbonyloxy)-1-azoniabicy
    clo [2.2.2] octane; bromide
30
    1-(2-Phenoxyethyl)-3(R)-(9[H]-xanthene-9-carbonyloxy)-1-azoniabic
    yclo [2.2.2]octane; bromide
    1-(3-Phenoxypropyl)-3(R)-(9[H]-xanthene-9-carbonyloxy)-1-azoniabi
    cyclo [2.2.2]octane; bromide
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1-Phenethyl-3(R)-(9[H]-xanthene-9-carbonyloxy)-1-azoniabicyclo[2.

2.2] octane; bromide

3(R)-(9-Hydroxy-9[H]-xanthene-9-carbonyloxy)-1-(3-phenoxypropyl)-

1- azoniabicyclo[2.2.2]octane; bromide

5 3(R)-(9-Hydroxy-9[H]-xanthene-9-carbonyloxy)-1-phenethyl-1-azonia bicy clo[2.2.2]octane; bromide

3(R)-(9-Hydroxy-9H-xanthene-9-carbonyloxy)-1-(3-thien-2-ylpropyl)

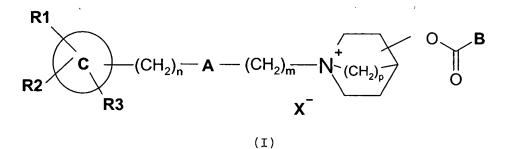
-1-azoniabicyclo[2.2.2]octane; bromide or

3(R)-(9-Methyl-9[H]-xanthene-9-carbonyloxy)-1-(3-phenoxy-propyl)-

10 1-azonia-bicyclo[2.2.2]octane; bromide

- 21. A compound according to any one of the preceding claims characterised in that it has an IC_{50} value for muscarinic M_3 receptors (Hm3) of less than 35 nM.
- $$\tt 22.$$ A process for the preparation of a compound of formula 15 (I)

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which comprises reacting an alkylating agent of formula (II)

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R1
$$(CH_2)_n$$
 A $(CH_2)_m$ X

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(II)

with a compound of formula (III)

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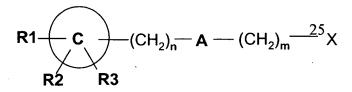
$$N_{(CH_2)_p}$$
 O B

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(III)

wherein, in each of formulae I, II and III, R^1 , R^2 , R^3 , \bigcirc , A, X, B, n, m and p are as defined in any one of claims 1 to 20.

- 23. A process according to claim 22 characterised in that 20 the resulting reaction mixture is purified by solid phase extraction.
 - 24. A compound of formula (II)



(II)

- 30 wherein R^1 , R^2 , R^3 , \bigcirc , A, X, n and m are as defined in any one of claims 1, 2, 4 to 11, 14 or 20.
 - 25. A compound of formula (III)

$$N_{(CH_2)_p}$$
 O B

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(III)

wherein B and p are as defined in any one of claims 1 to 3, 12, 13 or 15 to 20, and having the (R)-configuration.

- 26. A compound according to claim 25 which is 9-Methyl-9[H]-fluorene-9-carboxylic acid 1-azabicyclo[2.2.2]oct-3(R) -yl ester; 9-Methyl-9[H]-xanthene-9-carboxylic acid 1-azabicyclo[2.2.2]oct-3(R) -yl ester; 2-Hydroxy-2,2-difuran-2-yl-acetic acid -azabicyclo[2.2.2]oct-3(R) -yl ester.
 - 27. A compound of formula (VII)

(VII)

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wherein p and R^8 are as defined in any one of claims 1 to 3 or 25 12.

- 28. A compound according to claim 27, wherein \mathbb{R}^8 is a 2-thienyl or 2-furyl group.
- 29. A compound according to claim 27 which is Oxothien-2-yl-acetic acid 1-azabicyclo[2.2.2]oct-3(R)-yl ester; or Oxofuran-2-yl-acetic acid 1-azabicyclo[2.2.2]oct-3(R)-yl ester.
 - 30. Use of a compound according to any one of claims 24 to 29 in a process for producing a compound of formula (I) as defined in any one of claims 1 to 20.

- 31. A pharmaceutical composition comprising a compound according to any one of claims 1 to 21 in admixture with a pharmaceutically acceptable carrier or diluent.
- 32. A compound according to any one of claims 1 to 21, or 5 a pharmaceutical composition according to claim 31 for use in a method of treatment of the human or animal body by therapy.
 - 33. Use of a compound according to any one of claims 1 to 21, or a pharmaceutical composition according to claim 31 for the manufacture of a medicament for use in the treatment of respiratory, urinary or gastrointestinal disease.
 - 34. Use of a compound according to any one of claims 1 to 21 or a pharmaceutical composition according to claim 31 for the manufacture of a medicament for use in the treatment of COPD, chronic bronchitis, asthma and rhinitis.
- 35. A method for treating respiratory, urinary and/or gastrointestinal disease which method comprises administering to a human or animal patient in need of such treatment an effective amount of a compound according to any one of claims 1 to 21 or of a pharmaceutical composition according to claim 31.

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